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Uterine Cervix Metastasis as an Initial Manifestation of Metastatic Renal Cell Carcinoma: A Case Report and Review of Literature

Muhannad AlArifi¹, Khalid Riaz², Mutahir Ali Tunio^{2*} and Mushabbab Al Asiri³

- ¹Muhannad AlArifi, King Saud bin AbdulAziz University for Health Sciences, Riyadh 11345, Saudi Arabia
- ²Khalid Riaz, Radiation Oncology, Comprehensive Cancer Centre, King Fahad Medical City (KFMC), Riyadh 59046, Saudi Arabia
- ³Mushabbab Al Asiri, Director Comprehensive Cancer Center, Chairman Radiation Oncology, King Fahad Medical City, Riyadh-59046, Saudi Arabia

Abstract

Background: Renal cell carcinoma (RCC) has unpredictable and diverse behaviour. The classic triad of hematuria, loin pain and abdominal mass is uncommon at time of presentation. About 25%-30% of patients are found to have metastases at the time of diagnosis. Bones, lungs, liver and brain are the frequent sites of metastases. RCC with metastasis to the cervix uteri is rarest manifestation and only four case reports have been published so far.

Case Presentation: Herein we present a case of 70 year old Saudi female presenting with 4 months history of vaginal bleeding and weight loss. Her past medical history revealed left sided radical nephroectomy for RCC. She had no other co-morbidities. On physical examination, she was found emaciated and per vaginal examination showed fragile fungating mass of cervix. The punch biopsy of cervical mass confirmed the diagnosis of metastatic RCC. Further, staging workup showed bilateral pulmonary metastasis. She was given palliative Radiotherapy 30 Gy in 10 fractions followed by Sunitinib 50 mg oral daily, but patient died of progressive disease 4 months of palliative radiotherapy.

Conclusion: Metastatic RCC to cervix uteri is very rare manifestation. The physicians should consider metastasis from another primary as the differential diagnosis in order to plan optimal treatment. Reported treatment is radical hysterectomy with bilateral salpingo-oophorectomy followed by Tyrosine Kinase Inhibitors (TKIs). Patients who are not candidate for surgery; radiotherapy and TKIs is a reasonable option.

Keywords: Renal cell carcinoma; Metastasis; Cervix uteri; Rare manifestation

Introduction

Renal cell carcinoma (RCC) has unpredictable and diverse behavior. The incidence of RCC over last 20 years has progressively increased due to widespread use of modern imaging [1]. About 30%-50% of patients are found to have metastases at diagnosis. While bone, lymph nodes, lungs and brain constitute expected 'homing' sites, metastasis may turn up at the unusual locations (skin, testis, maxillary antrum and tongue) [2-4].

Cervix Uteri is rare site of metastasis from other primaries, because of dense stroma and less vascularity. Metastatic RCC to cervix uteri is extremely rare and usually associated with left sided RCC; only four case reports have been published so far [5-8]. The exact mechanism is not known; however retrograde venous flow of tumor cells from left renal vein to the left ovarian vein and cervical and vaginal venous plexus explains the spread of left sided RCC to cervix uteri [9].

Here we present a case of postmenopausal lady with cervix uteri metastasis as initial metastatic manifestation of RCC.

Case Presentation

A 70 year old Saudi female presented in our clinic with vaginal bleeding and significant weight loss. She had noticed vaginal bleeding for 4 months and it had been rapidly progressing over 2 weeks. Her previous surgical history revealed left sided radical nephrectomy four years back for RCC (Figure 1).

On physical examination, her vitals were stable. She was found severely malnourished and emaciated. There was no palpable lymphadenopathy and examination of chest, heart, nervous system and abdomen was normal. Per vaginal examination showed a fragile easily

bleeding fungating lesion of size 5×5 cm in exo-cervix with involvement of vaginal fornices and right parametrium.



Figure 1: Computed tomography (CT) of abdomen showing mass in left kidney for which radical nephrectomy was performed four years back.

*Corresponding author: Mutahir Tunio, Assistant Consultant, Radiation Oncology, Comprehensive Cancer Centre, King Fahad Medical City (KFMC), Riyadh 59046, Saudi Arabia, Tel: +966 1 2889999; Fax: 966 1 4614006; E-mail: drmutahirtonio@hotmail.com

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Magnetic resonance imaging (MRI) of pelvis showed necrotic mass of cervix uteri mass of size 6×5.5 cm with bilateral parametrial invasion and no pelvic lymphadenopathy (Figure 2). Computed tomography (CT) abdomen showed absent left kidney consistent with left radical nephrectomy for RCC with no recurrence in the renal bed and CT chest revealed bilateral pulmonary metastasis (Figure 3). Serum chemistry, hematology and bone scan were found normal.

Punch biopsy of mass of cervix uteri was performed, which revealed clear cell histology consistent with metastatic RCC (Figure 4).

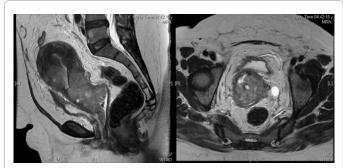


Figure 2: Magnetic resonance imaging (MRI) of pelvis showing 5 x 5 m cervical mass with parametrial invasion.

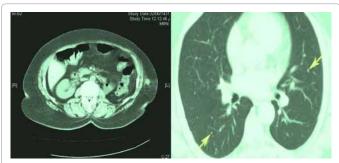


Figure 3: Computed tomography (CT) of (a) abdomen showing left nephrectomy renal bed with no recurrence and (b) chest showing pulmonary metastasis

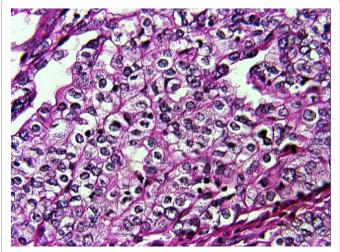


Figure 4: Histopathology of cervical mass showing clear cell consistent with renal cell carcinoma.

Author	Age and presenting complaints	Treatment offered	Survival
Seseke F, et al. [5]	Postmenopausal Vaginal bleeding	Total abdominal hysterectomy and bilateral salpingo-oophorectomy	NA
Bozaci EA, et al. [6]	19 years Vaginal bleeding	Total abdominal hysterectomy and bilateral salpingo-oophorectomy	NA
Zafrakas M, et al. [7]	45 years Vaginal bleeding	Total abdominal hysterectomy and bilateral salpingo-oophorectomy and pelvic lymph node dissection And Sunitinib 50 mg oral daily	NA
Godfrey GJ, et al. [8]	Postmenopausal lady Cervical polyp	Total abdominal hysterectomy and bilateral salpingo-oophorectomy	NA
Our case	70 years Vaginal bleeding and weight loss	Palliative radiotherapy and Sunitinib 50 mg oral daily	4 months

Table 1: Previously published case reports on Renal Cell Carcinoma metastasizing to cervix uteri.

After presenting this case in a multidisciplinary tumor (MDT) meeting, patient was given external beam radiation therapy (EBRT) 30 Grays (Gy) in 10 fractions to cervix uteri mass to control per vaginal bleeding. Post radiation therapy, her vaginal bleeding was much controlled and she was sent to medical oncology team and patient was started on sunitinib 50 mg oral daily. Patient died of progressive pulmonary disease 4 months of palliative EBRT.

Discussion

Relatively few case reports of primary extra-pelvic carcinomas with metastasis to cervix uteri have been published. The most frequent primary carcinomas metastasizing to uterus and cervix are the breast, stomach, ovarian and colorectal cancers [10].

Renal Cell carcinoma (RCC) may remain clinically occult for the most of its course and the tumor in the kidney may progress unnoticed to a large mass until metastases appear. About two third of cases of RCC metastasize to lungs, liver and bones. Only less than 0.5% cases metastasize to female genital system [9]. Uterine cervix metastasis as an initial manifestation of metastatic RCC is extremely rare and only four cases have been published so far Table 1. All case reported had left sided RCC, which explains the possible route of spread of renal tumor cells via retrograde venous flow from left renal vein, left ovarian vein and then cervical venous plexus [7]. The treatment of choice for uterine cervix metastasis is the surgical excision in form of total abdominal hysterectomy and bilateral salpingo-oophorectomy (TAH+BSO) with or without lymph node dissection followed by TKIs such Sunitinib, Sorafenib and Temsirolimus.

Our patient was not surgical candidate because of extensive lung metastases. We achieved good haemostatic control of vaginal bleeding by palliative radiotherapy. Prognosis of uterine cervix metastasis depends on presence of metachronous metastasis as our patient died within four months of treatment secondary to lung metastasis.

Conclusion

Metastatic RCC to cervix uteri is very rare manifestation. The physicians should consider metastasis from another primary as the differential diagnosis in order to plan optimal treatment. Reported treatment is TAH+BSO followed by TKIs. Patients who are not candidates for surgery; radiotherapy and TKIs are a reasonable option.

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